

"Il" (") (") (") Alm Alm (") (") (") (") (") (") (") (")

"H" H"H Alles H"b all. H"b" Ber Hall

Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE

ntors: Linda SHERMAN and Joseph LUSTGAPTEN ication No.: February 1, 2001 Docket No.: 313332000101

eet 1 of 17

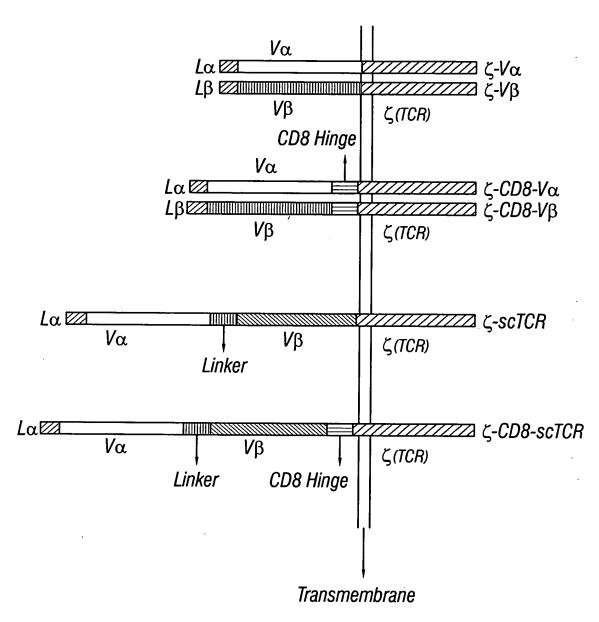


FIG.1

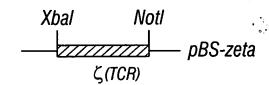
<u>|</u>=1

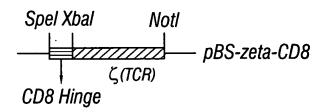
Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE ntors: Linda SHERMAN and Joseph LUSTGADEN

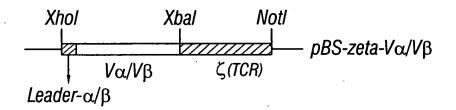
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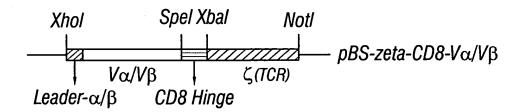
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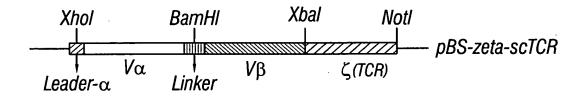
Smeet 2 of 17











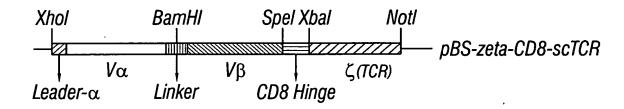


FIG. 2

"#" 6"% "IL., 6""! 41", 4"" u Ir 4...p 11 9...s 4...s 4...s 4...s

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Docket No.: 313332000101

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		HIG																ίσεΩ	ענ	WO.	1]
L	E	M	Q	R	N	L	G	A	V	L	G	I	L	W	V	Q	I	[SEQ	ID	NO:	2]
		63			72			81			90			99			108				
TGC	TGG	CTG															CAG				
	 W	 L		 E	0	 Q		0	0	 S			 S	 T.		 T.	0				
Ů	•	_													•	-	¥				
CAC	ccc	117				СШС										CAC					
GAG		GAG	AAC	GCA	GAG		CAG	TGT	AGC	TTT	TUU	ATC	TIT	ACA	AAC	CAG					
E	G <sub>.</sub>	E	N	A	E	L	Q	C	S	F	S	Ι	F	T	N	Q	V				
		171			180			189			198			207			216				
CAG	TGG	TTT.																			
								+	11011	CIC	010	NUC				M	CCI				
Λ																					
Q	W	 F	 Ү	Q	 R	 P					 V				 Y	 N	 P				
-		225		~	234		 G	 G 243	 R	 L	V 252	<b>S</b>	 L	L 261	 Y	 N	P 270				
-				~	234		 G	 G 243	 R	 L	V 252	<b>S</b>	 L	L 261	 Y	 N	P 270				
TCT		225 ACA	AAG	CAG	234 AGT		G AGA	G 243 CTG	R ACA	L TCC	V 252 ACA	S ACA	L GTC	L 261 ATT	 У ААА	N GAA	P 270 CGT				
TCT	GGG	225 ACA  T	AAG  K	CAG  Q	234 AGT  S	GGG  G	G AGA	G 243 CTG L	R ACA T	TCC	V 252 ACA  T	S ACA	L GTC  V	261 ATT  I	 Y ААА  K	N GAA  E	P 270 CGT R				
TCT  S	GGG  G	225 ACA	AAG  K	CAG  Q	234 AGT  S 288	GGG  G	G AGA  R	G 243 CTG L 297	R ACA T	TCC	V 252 ACA T 306	S ACA	L GTC  V	261 ATT  I 315	 Ү ААА  К	N GAA  E	P 270 CGT  R 324				

FIG. 3A

APPROYED O.G. FIG. CLASS SUBCLASS ΒÝ DRAFTSMAN

"H" ("I the start of the start

"F" 8"1 "L. 6"4 45" 4"1 10

Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE

Inventors: Linda SHERMAN and Joseph LUSTGAPTEN lication No.: February 1, 2001
Secket No.: 313332000101
Sneet 4

neet 4 of 17

4/17

																•	⊀
TGT	GCC	333 TCA		TCT			AGC			AAG	360 CTA	ACC		369 GGG		GGC	378 ACT
C	 A	S	N	S	G	G	S	N	A	K	L	T	F	G	K	G	T
AAA	CTC	387 TCT	GTT	AAA							414 GGC					GGG	
K	L	S	V	K	S	G	G	G	G	S	G	G	G	G	S	G	G
GGA	GGC	441 TCA									468 AGA					GTA	486 ACA
G	G	S	E	A	A	V	T	Q	S	P	R	N	K	V	A	V	T
GGA	GGA	495 AAG									522 AAT						
G	G	K	V	 T	L	S	C	N	Q	T	N	N	H	N	N	M	Y
TGG	TAT	549 CGG									576 CTG			585 TAT		TAT	594 GGT
W	Y	R	Q	D	T	G	Н	G	L	R	L	I	Н	Y	S	Y	G
GCT	GGC	603 AGC									630 GGA						
 A	G	\$	 T	 E	 K	G	D	I	P	D	G	Y	K	A	S	R	P

FIG. 3B

Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE Inventors: Linda SHERMAN and Joseph LUSTGAPTEN

ication No.: February 1, 2001 bocket No.: 313332000101

Smeet 5 of 17

5/17

AGC	CAA	657 GAG	AAC	TTC 	666 TCC	CTC	ATT	675 CTG	GAG	TTG 	684 GCT	ACC		693 TCT			702 TCA
S	Q	E	N	F	S	L	Q	L	E	L	A	T	P	S	I	T	S
GTG	TAC	711 TTC	TGT	GCC	720 AGC	GGT	GAG	729 ACA	GGG	ACC		GAA		747 TTA	TTT	TTC	756 GGT
V	<u></u>	F	C	A	S	G	E	T	G	T	N .	E	R	L	F	F	· G
CAT	GGA	765 ACC	AAG	CTG	774 TCT	GTC	CTG	783 ACT	AGT	AAC	792 TCC	ATC	ATG	801 TAC	TTC	AGC	810 CAC
Н	 G.	 T	 K		S	V	L	T	S	N	S	Ι	M	Y	F	S	H
TTC	GTG	819 CCG	GTC	TTC	828 CTG	CCA	GCG	837 AAG	CCC	ACC	846 ACG	ACC	G CCA	855 GCG	CCG	CGA	864 CCA
F	V	 P	V	F	L	P	A	K	P	· Т	T	T	P	A	P	R	P
CCA	A ACI	873 A CC0	} G GC(	G CCC	882 ACC	: ATC	C GCG	891 TCG	CA(	G CC	90( C CT(	) G TC(	C CT(	909 G CG(	) C CC <i>I</i>	A TCT	918 AGT
- <u>-</u> -	 T	 P	 A	P	T	I	A	. <b></b> -	Q	P	L	S	L	R	P	S	S
TC	T AG	92' A GA'	7 T CC	C AA	936 A CT(	6 C TG(	C TA	945 C CT(	o G CT	G GA	95 T GG	4 A AT	C CT				972 I GGT
 S	 R	 D	 P	 K	 L	 C	Y		L	 D	 G	 I	 L	F	I	Y	G

FIG. 3C

APPROVED O.G. FIG.

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Tr. 6"4 ... 6"1 65". 6"4 10

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lication No.: February 1, 2001 Docket No.: 313332000101

Sneet 6 of 17

6/17

																: -	
GTC	ATT	981 CTC	ACT	GCC	990 TTG	TTC	CTG	999 AGA	GTG	1 AAG	.008 TTC	AGC	1 AGG	.017 AGC	GCA	GAC	026 GČC :
٧	 I	L		 A	<u>r</u>	 F	L	R	V	<u>K</u>	F	S	R	S	Α	D	A
CCC	GCG	1035 TAC	CAG	1 CAG	.044 GGC	CAG	AAC	.053 CAG	CTC	TAT	1062 AAC	GAG	CTC	1071 AAT	CTA	GGA	.080 CGA
P	A	Y	Q	. Q	G	Q	N	Q	L	Y	N	E	L	N	L	G	R
AGA				GAT													134 GGG
 R	<u></u>	 E	Y	D	V	L	D	K	R	R	G	R	D	P	E	M	G
GGA	AAG	1143 CCG		AGG													1188 AAA
G	 K	P	R	R	K	N	P	Q	E	G	L	Y	N	E	L	Q	K
GAT	AAG	1197 ATG	GCG	GAG	1206 GCC	TAC	AGT	1215 GAG	ATT	GGG	1224 ATG	AAA	GGC	1233 GAG	CGC	CGG	1242 AGG
D	K	 M	A	E	A	Y	S	E	I	G	M	K	G	E	R	R	R
GGC		1251 GGG	CAC	GAT	1260 GGC	CTT	TAC	CAG	GGT	CTC	AGT	ACA	GCC	1287 : ACC	. AAG	GAC	1296 ACC
G	K	 G	Н	D	G	L							A	T	K	D	T
TAC	GAC	1305 : GCC	i CTI	CAC	1314 : ATG	CAG	GCC	1323 CTC	} G CCC	CCT	1332 CGC	? C TAA			C GCC		1350 GCG
Y	D		 L	Н	M	Q	A	L	P	P	R	*	A	A	A	T	A

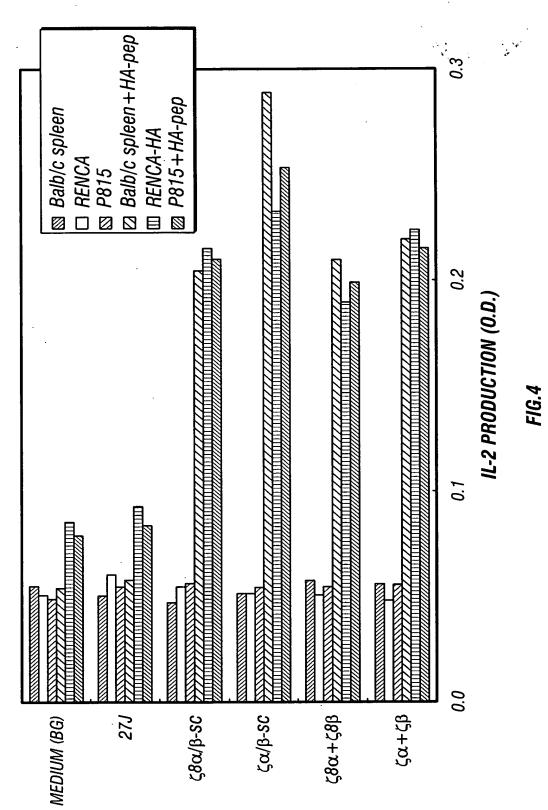
FIG.3D

APPROVED O.G. FIG.
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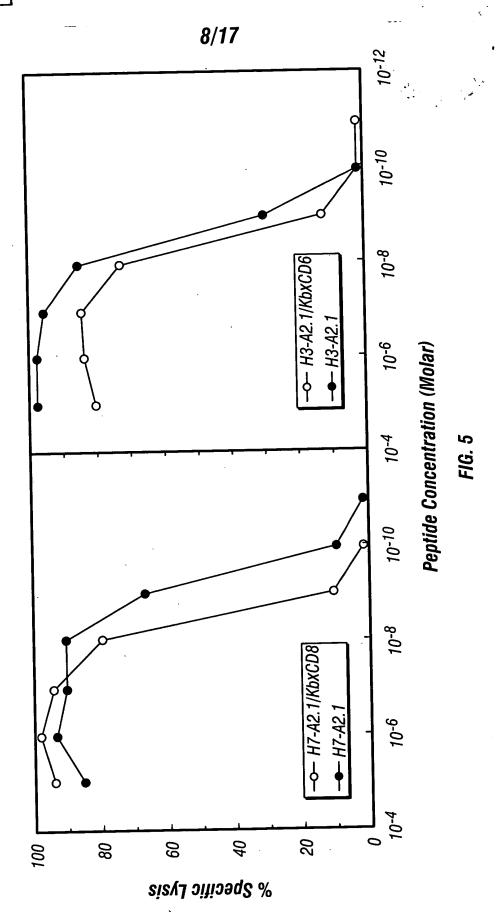
Sneet 7 of 17



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APPROVED O.G. FIG.

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19" 4"19 offen 21"19 16"11. 16"11 10.

Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE Inventors: Linda SHERMAN and Joseph LUSTGARTEN

cation No.: February 1, 2001 Docket No.: 313332000101

Smeet 9 of 17

9/17

## Alpha Groups

Vα1 Vα2 Vα3 Vα4.1,2,3 Vα4.4	TGA GACTG CATCC CO	AG GCA AC AAA AG CTG GG AGA CA GCA	GTC CTC AGG	CCC CTC TCC	AAT AAG ACA	CTC TAC GTT	TGA TAT CCT	CAG TC CTT	T	[SEQ [SEQ [SEQ [SEQ	ID ID ID	NO: NO: NO:	4] 5] 6]
2. Va5 Va6 Va7 Va8 Va9	GGT TO GCG A GAG C	GG TCT CC TCT AG AAC TC CAC TG CGA	TCA TCA AGA	GGG CCC CAA	TCC TGG CAA	AGA ACT GAG	ATA GTT GAC	TGT CAT GCA		[SEQ [SEQ [SEQ [SEQ	ID ID ID	NO: NO: NO:	9] 10] 11]
3. Va10 Va11 Va12 Va14 Va15	CAG C CCA C ACC T	GT CAG CT CAT CCA GGG CG AGA	CAA ACC GAA	TCT ACA TCC	GTT GTT TAA	CTA TAT GCT	CTT CAT CAT	GGC TCA CAT	T A	[SEQ [SEQ [SEQ	ID ID ID	NO: NO: NO:	14] 15] 16]
4. Vα16 Vα17 Vα18 Vα19 Vα20	CAA ( GTG 1 CCG 1	GCA AAC GAC ATC FAT GAA FAT TTC AGC TC	CAT A ACC TTT	AAC CAG CTT	TGC GAC ATG	CCT AGT TTG	ACA TCT TTT	G TAC TGG	AT	[SEQ [SEQ [SEQ	ID ID ID	NO: NO: NO:	18] 19] 20] 21] 22]

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Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE

Inventors: Linda SHERMAN and Joseph LUSTGARTEN lication No.: February 1, 2001

lication No.: February 1, 2 bocket No.: 313332000101 Sheet 10 of 17

	Beta Groups													
	1. Vβ1 Vβ2 Vβ3 Vβ4 Vβ5.1	GGC GTG GAT		TGG GGG CGA	TAC CAA ACA	CAC GGA GTA	GTG CAA TCT	GTC AAA AGG	AA GC C	•	[SEQ [SEQ [SEQ [SEQ	ID ID ID	NO: NO: NO:	<ul><li>24]</li><li>25]</li><li>26]</li></ul>
1	2. Vβ6 Vβ7 Vβ8.1 Vβ8.2 Vβ8.3	TAC ATA ATA	TGA CTG ACC ACC GCC	ATC ATG ACA	AAA ACA ACA	AGA ATA ACA	ATG TGT TGT	GGA ACT ACT	GA GG GG		[SEQ [SEQ [SEQ [SEQ	ID ID ID	NO: NO: NO:	29] 30] 31]
ייניים ייניים איניים לעיים מיינים איניים איניים איניים ייניים איניים	3. Vβ9 Vβ10 Vβ11 Vβ12 Vβ13	GAT ACA ACC	TTG TAT AGG TAC TTG	GTT TGA AGA	TAG CAG ACC	CTA GGA CAA	CAA AGG GGA	TAA GAC CTC	TA AA AG		[SEQ [SEQ [SEQ [SEQ [SEQ	ID ID ID	NO: NO: NO:	34] 35] 36]
	4. Vβ14 Vβ15 Vβ16 Vβ17 Vβ18	AGA CAT GTA	GAG ACC CAA GTC CTG	ATC ATA CTG	TGT ATA AAA	AAG GAT AAG	AGT ATG GGC	GGA GGG ACA	AC CA CT		[SEQ [SEQ [SEQ [SEQ	ID ID ID	NO:	39] 40] 41]

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18" #"14 dlas 4"11 sil. 4"14 il.

Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE gentors: Linda SHERMAN and Joseph LUST?

plication No.: February 1, 2001

Docket No.: 313332000101

Sheet 11 of 17

11/17

45 18 27 36 ATG AAA TCC TTG AGT GTT TCC CTA GTG GTC CTG TGG CTC CAG TTA AAC TGG GTG [SEQ ID NO: 43] Met Lys Ser Leu Ser Val Ser Leu Val Val Leu Trp Leu Gln Leu Asn Trp Val [SEQ ID NO: 44] 90 99 108 63 72 81 CAG AGC CAG CAG AAG GTG CAG CAG AGC CCA GAA TCC CTC AGT GTC CCA GAG GGA Gln Ser Gln Gln Lys Val Gln Gln Ser Pro Glu Ser Leu Ser Val Pro Glu Gly 153 162 135 144 126 117 GGC ATG GCC TCT CTC ACC TGC ACT TCA AGT GAT CGC AAT TTT CAG TAT TTC TGG --- --- --- --- --- --- --- --- --- --- --- --- ---Gly Met Ala Ser Leu Asn Cys Thr Ser Ser Asp Arg Asn Phe Gln Tyr Phe Trp 207 189 198 216 180 171 TGG TAC AGA CAG CAT TCT GGA GAA GGC CCC AAA GCA CTG ATG TCC ATC TTC TGG Trp Tyr Arg Gln His Ser Gly Glu Gly Pro Lys Ala Leu Met Ser Ile Phe Ser

FIG. 7A

4..... 11.... 14.... 14.

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Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE ntors: Linda SHERMAN and Joseph LUSTG

ication No.: February 1, 2001 Docket No.: 313332000101

Sheet 12 of 17

12/17

261 270 252 225 234 243 GAT GGT GAC AAG AAA GAA GGC AGA TTC ACA GCT CAC CTC AAT AAG GCC AGC CTG Asp Gly Asp Lys Lys Glu Gly Arg Phe Thr Ala His Leu Asn Lys Ala Ser Leu 306 315 324 279 288 297 CAT GTT TCC CTG CAC ATC AGA GAC TCC CAG CCC AGT GAC TCC GCT CTC TAC TTC His Val Ser Leu His Ile Arg Asp Ser Gln Pro Ser Asp Ser Ala Leu Tyr Phe 369 360 378 351 333 . 342 TGT GCA GTT ATG GAT TAT AAC CAG GGG AAG CTT ATC TTT GGG CAG GGT ACC AAG Cys Ala Val Met Asp Tyr Asn Gln Gly Lys Leu Ile Phe Gly Gln Gly Thr Lys 387 TTA TCT ATC AAG CCC 3' Leu Ser Ile Lys Pro

FIG. 7B

Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE Linda SHERMAN and Joseph LUSTGA N

Cation No.: February 1, 2001 Docket No.: 313332000101

Sheet 13 of 17

13/17

45 18 27 36 54 ATG GGC TCC AGA CTC TTC TTT GTG GTT TTG ATT CTC ATG TGT GCA AAA CAC ATG [SEQ ID NO: 45] Met Gly Ser Arg Leu Phe Phe Val Val Leu Ile Leu Cys Ala Lys His Met [SEQ ID NO: 46] 63 72 99 81 90 108 GAG GCT GCA GTC ACC CAA AGT CCA AGA AGC AAG GTG GCA GTA ACA GGA GGA AAG --- --- --- --- --- --- --- --- --- --- --- --- --- ---Glu Ala Ala Val Thr Gln Ser Pro Arg Ser Lys Val Ala Val Thr Gly Gly Lys 117 126 153 135 144 162 GTG ACA TTG AGC TGT CAC CAG ACT AAT AAC CAT GAC TAT ATG TAC TGG TAT CGG --- --- --- --- --- --- --- --- --- --- --- --- --- ---Val Thr Leu Ser Cys His Gln Thr Asn Asn His Asp Tyr Met Tyr Trp Tyr Arg 171 180 189 198 207 216 CAG GAC ACG GGG CAT GGG CTG AGG CTG ATC CAT TAC TCA TAT GTC GCT GAC AGC Gln Asp Thr Gly His Gly Leu Arg Leu Ile His Tyr Ser Tyr Val Ala Asp Ser

FIG. 7C

"H" 6"H 1,H2, 6" t 162, 6" t 10

Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE ntors: Linda SHERMAN and Joseph LUSTGATEN

leation No.: February 1, 2001 Docket No.: 313332000101

Sheet 14 of 17

14/17

		225			234			243			252			261			270
ACG	GAG	AAA	GGA	GAT	ATC	CCT	GAT	GGG	TAC	AAG	GCC	TCC	AGA	CCA	AGC	CAA	GAG
Thr	Glu	Lys	Gly	Asp	Ile	Pro	Asp	Gly	Tyr	Lys	Ala	Ser	Arg	Pro	Ser	Gln	Glu
		220			200			207			20.0	•		21 [			204
														315			324
AAT	TTC	TCT	CTC	ATT	CTG	GAG	TTG	GCT	TCC	CTT	TCT	CAG	TCA	GCT	GTA	TAT	TTC
Asn	Phe	Ser	Leu	Ile	Leu	Glu	Leu	Ala	Ser	Leu	Ser	Gln	Ser	Ala	Val	Tyr	Phe
		333			342			351			360			369			378
TGT	GCC	AGC	AGC	GAT	TTC	GCC	GGG	ACA	GGG	GGC	TTC	TAT	GAA	CAG	TAC	TTC	GGT
	<del></del>																
Cys	Ala	Ser	Ser	Asp	Phe	Ala	Gly	Thr	Gly	Gly	Phe	Tyr	Glu	Gln	Tyr	Phe	Gly
		387			396								•				
000																	
	GGC	ACC	AGG	$\mathtt{CTC}$	ACG	GTT	TCT	31									
					ACG		TCT	31									

FIG. 7D

"H" H"H ..H., G"H AT., H"H 10 10- q.,P H H 40, H, Hp. H Title: T CELL RECEPTORS FROM HLA TRANSGENIC MICE antors: Linda SHERMAN and Joseph LUSTG

lication No.: February 1, 2001 Docket No.: 313332000101

Sheet 15 of 17

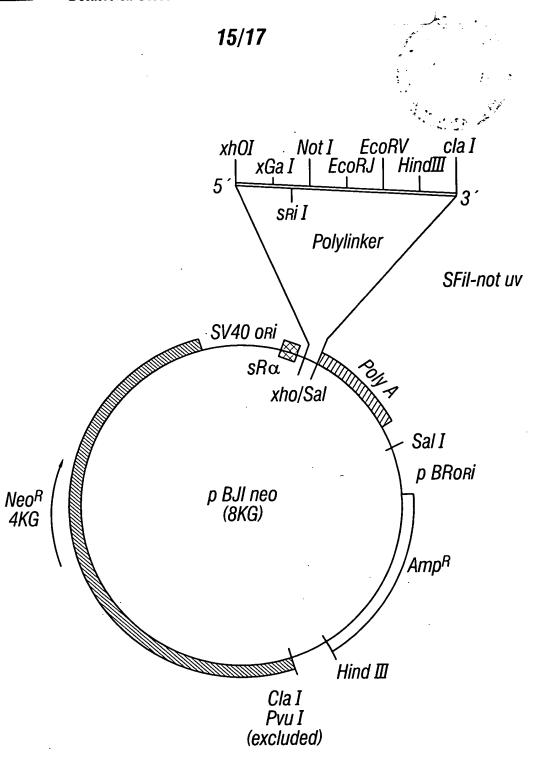


FIG. 8

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Docket No.: 313332000101

Sheet 16 of 17

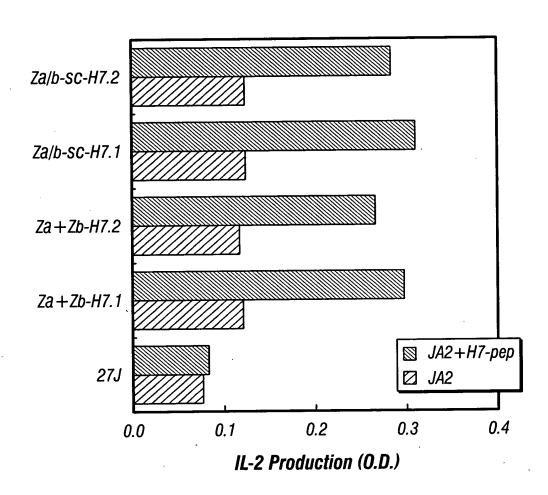


FIG. 9

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cation No.: February 1, 2001 Docket No.: 313332000101

Sheet 17 of 17

17/17

